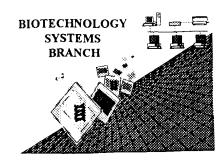
12/49

# RAW SEQUENCE LISTING ERROR REPORT



The Biotechnology Systems Branch of the Scientific and Technical Information Center (STIC) detected errors when processing the following computer readable form:

09/893,666	
0196	
12/6/2001	
	1 /

THE ATTACHED PRINTOUT EXPLAINS DETECTED ERRORS.
PLEASE FORWARD THIS INFORMATION TO THE APPLICANT BY EITHER:

- 1) INCLUDING A COPY OF THIS PRINTOUT IN YOUR NEXT COMMUNICATION TO THE APPLICANT, WITH A NOTICE TO COMPLY or,
- 2) TELEPHONING APPLICANT AND FAXING A COPY OF THIS PRINTOUT, WITH A NOTICE TO COMPLY FOR CRF SUBMISSION QUESTIONS, PLEASE CONTACT MARK SPENCER, 703-308-4212.

FOR SEQUENCE RULES INTERPRETATION, PLEASE CONTACT ROBERT WAX, 703-308-4216. PATENTIN 2.1 e-mail help: patin21help@uspto.gov or phone 703-306-4119 (R. Wax) PATENTIN 3.0 e-mail help: patin3help@uspto.gov or phone 703-306-4119 (R. Wax)

TO REDUCE ERRORED SEQUENCE LISTINGS, PLEASE USE THE CHECKER VERSION 3.0 PROGRAM, ACCESSIBLE THROUGH THE U.S. PATENT AND TRADEMARK OFFICE WEBSITE. SEE BELOW:

## Checker Version 3.0

The Checker Version 3.0 application is a state-of the-art Windows based software program employing a logical and intuitive user-interface to check whether a sequence listing is in compliance with format and content rules. Checker Version 3.0 works for sequence listings generated for the original version of 37 CFR §§1.821 – 1.825 effective October 1, 1990 (old rules) and the revised version (new rules) effective July 1, 1998 as well as World Intellectual Property Organization (WIPO) Standard ST.25.

Checker Version 3.0 replaces the previous DOS-based version of Checker, and is Y2K-compliant. Checker allows public users to check sequence listings in Computer Readable form (CRF) before submitting them to the United States Patent and Trademark Office (USPTO). Use of Checker prior to filing the sequence listing is expected to result in fewer errored sequence listings, thus saving time and money.

Checker Version 3.0 can be down loaded from the USPTO website at the following address: http://www.uspto.gov/web/offices/pac/checker

### Raw Sequence Listing Error Summary

ERROR DETECTED	suggested correction serial number: $09/893,666$
ATTN: NEW RULES CASES	S: PLEASE DISREGARD ENGLISH "ALPHA" HEADERS, WHICH WERE INSERTED BY PTO SOFTWAL
1 Wrapped Nucleics Wrapped Aminos	The number/text at the end of each line "wrapped" down to the next line. This may occur if your file was retrieved in a word processor after creating it. Please adjust your right margin to .3; this will prevent "wrapping."
2Invalid Line Length	The rules require that a line not exceed 72 characters in length. This includes white spaces.
3Misaligned Amino Numbering	The numbering under each 5th amino acid is misaligned. Do not use tab codes between numbers; use space characters, instead.
4Non-ASCII	The submitted file was not saved in ASCII(DOS) text, as required by the Sequence Rules. Please ensure your subsequent submission is saved in ASCII text.
5Variable Length	Sequence(s) contain n's or Xaa's representing more than one residue. Per Sequence Rules, each n or Xaa can only represent a single residue. Please present the maximum number of each residue having variable length and indicate in the <220>-<223> section that some may be missing.
6PatentIn 2.0 "bug"	A "bug" in PatentIn version 2.0 has caused the <220>-<223> section to be missing from amino acid sequences(s)  Normally, PatentIn would automatically generate this section from the previously coded nucleic acid sequence. Please manually copy the relevant <220>-<223> section to the subsequent amino acid sequence. This applies to the mandatory <220>-<223> sections for Artificial or Unknown sequences.
7Skipped Sequences (OLD RULES)	Sequence(s) missing. If intentional, please insert the following lines for each skipped sequence: (2) INFORMATION FOR SEQ ID NO:X: (insert SEQ ID NO where "X" is shown) (i) SEQUENCE CHARACTERISTICS: (Do not insert any subheadings under this heading) (xi) SEQUENCE DESCRIPTION:SEQ ID NO:X: (insert SEQ ID NO where "X" is shown) This sequence is intentionally skipped
	Please also adjust the "(ii) NUMBER OF SEQUENCES:" response to Include the skipped sequences.
8 Skipped Sequences (NEW RULES)	Sequence(s) missing. If Intentional, please insert the following lines for each skipped sequence. <210> sequence id number <400> sequence id number 000
9Use of n's or Xaa's (NEW RULES)	Use of n's and/or Xaa's have been detected in the Sequence Listing.  Per 1.823 of Sequence Rules, use of <220>-<223> is MANDATORY if n's or Xaa's are present.  In <220> to <223> section, please explain location of n or Xaa, and which residue n or Xaa represents.
0 V Invalid <213> Response	Per 1.823 of Sequence Rules, the only valid <213> responses are: Unknown, Artificial Sequence, or scientific name (Genus/species). <220>-<223> section is required when <213> response is Unknown or is Artificial Sequence
1Use of <220>	Sequence(s) missing the <220> "Feature" and associated numeric identifiers and responses.  Use of <220> to <223> is MANDATORY if <213> "Organism" response is "Artificial Sequence" or "Unknown." Please explain source of genetic material in <220> to <223> section.  (See "Federal Register," 06/01/1998, Vol. 63, No. 104, pp. 29631-32) (Sec. 1.823 of Sequence Rules)
2Patentin 2.0 "bug"	Please do not use "Copy to Disk" function of PatentIn version 2.0. This causes a corrupted file, resulting in missing mandatory numeric identifiers and responses (as indicated on raw sequence listing). Instead, please use "File Manager" or any other manual means to copy file to floppy disk.
3Misuse of n	n can only be used to represent a single nucleotide in a nucleic acid sequence. N is not used to represent any value not specifically a nucleotide.

AMC/MH - Biotechnology Systems Branch - 08/21/2001

OIPE

RAW SEQUENCE LISTING

DATE: 12/06/2001 TIME: 10:36:14

PATENT APPLICATION: US/09/893,666

Input Set : A:\210217US0.ST25.txt

5	<110 <120	> T	ITLE	OF I	INVE	OITE	<b>ч:</b> н:	igh e	estr	-		siti <sup>.</sup>	ve me	edaka	a fis	sh	
	<pre>1 :130 &gt; FILE REFERENCE: 210217US-620-7249-0 1 &lt;140 &gt; CURRENT APPLICATION NUMBER: US 09/893,666</pre>																
	141> CURRENT APPLICATION NOMBER. 05 09/093/000																
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	-160																
	.:170					entIi	n ve:	rsio	n 3.	L						Not	compi'
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	qcc		ccc	aac	aat.	gaa		act.	ccc	ctt	tac		caq	tcc	agc	acc	330
	Ala																
	5 ئە					30					35					40	
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	${\tt Gl}_T^{\omega}$	Tyr	Tyr	Ser		Pro	Leu	Glu	Thr		Gly	Pro	Pro	Ser		Gly	
51	t	~+~		t a a	45 ata	G (7.2	3.7±	aaa	000	50	200	aat	ata	ata	55 +++	ata	426
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	ccc	tcc	agc		aga	ctc	agt	CCC	ttt	atg	cat	cca	CCC	agc	cac	cac	474
5.8	Pro	Ser	Ser	Pro	Arg	Leu	Ser	Pro	Phe	Met	His	Pro	Pro	Ser	His	His	
59			75					80					85				
	tat																522
62 63	Tyr	Leu 90	Glu	Thr	Thr	ser	Thr 95	Pro	val	Tyr	Arg	Ser 100	ser	Hls	GIN	GTA	
	gec		agg	aaa	gac	сас	, ,	aac	tee	caa	aaa		aco	tac	age	cta	570
	Ala																2,0
	105				- I.	110	<i>1</i> -	- 1		,	115			4		120	
	ggq																618
7()	Gly	Glu	Leu	Gly	Ala	Gly	Ala	Gly	Ala	Gly	Gly	Phe	Glu	Met	Ala	Lys	

DATE: 12/06/2001 PATENT APPLICATION: US/09/893,666 TIME: 10:36:14

Input Set : A:\210217US0.ST25.txt

71					125					130					135		
73	gac	acg	cgt.	ttc	tgc	gcc	gtg	tgc	agc	gac	tac	gcc	tct	ggg	tac	cac	666
						Āla											
75	_			140				_	145					150			
77	tat	ggg	gtg	tgg	tct	tgt	gag	ggc	tgc	aag	gcc	ttc	ttc	aag	agg	agc	714
78	Tyr	Gly	Val	Trp	Ser	Cys	Glu	Gly	Cys	Lys	Ala	Phe	Phe	Lys	Arg	Ser	
79			155					160					165				
81	atc	cag	ggt	cac	aat	gac	tat	atg	tgc	cca	gcg	acc	aat	cag	tgc	act	762
82	Ile	Gln	Gly	His	Asn	Asp	Tyr	Met	Cys	Pro	Ala	Thr	Asn	Gln	Cys	Thr	
83		170					175					180					
85	att	gac	aga	aat	cgg	agg	aag	agc	tgc	cag	gct	tgt	cgt	ctt	agg	aag	810
86	Ile	Asp	Arg	Asn	Arg	Arg	Lys	Ser	Cys	Gln	Ala	Cys	Arg	Leu	Arg	Lys	
87	185					190					195					200	
	_		-			atg	-					-					858
91)	Cys	Tyr	Glu	Val	Gly	Met	Met	Lys	Gly	Gly	Val	Arg	Lys	Asp	Arg	Ile	
91					205					210					215		
					_	gac						-		-		-	906
	Arg	Ile	Leu		Arg	Asp	Lys	Arg		Thr	Gly	Val	Gly		Gly	Asp	
95				220					225					230			
	-	-	-	_		cag				_				-			954
	Lys	Val		Lys	Gly	Gln	Glu		Lys	Thr	Val	His	-	Asp	Gly	Arg	
99			235					240					245				
						ı gga											1002
	_	_		Ser	Thr	Gly	_		, GT	/ Gly	7 Gly			GI	Arg	Leu	
103		250					255					260					1050
				-		cct			_		_				-		1050
			. Thr	ser	. 116	Pro		GIU	GIR	ı vaı	. Leu 275		ı Let	Let	i Gin	_	
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						ctc											1090
111		וובני.	PIC	PIC	285	Leu	Cys	ser	AIG	290		ь тес	i sei	. AIG	295		
		u a a a	ata	300		, ratg	200	ata	cto			, ato	race				1146
						Met											1140
115		'JI'U	. vai	300		. Met	1111	ьеи	305		Ser	nec	. Ald	310	_	GLU	
		ato	cac			gcc	taa	acc			cto	cca	uat			cad	1194
	-	_				Ala											11.71
119		, vai	315		. 110	. ma	112	320	_	, дуз	, пса		325		. neu	3111	
		tac			gat	. cag	ata			r cta	ааа	ago			r eta	gag	1242
	_		_		-	Gln		_	-	-		-	_		_		
123		330				0 2 1 1	335	200				340					
				atc	aac	ctc		t.aa	aga	r t.c.a	ato			. ccc	aaa	aaq	1290
						Leu											
	345				- 1	350		- <b>r</b> -	)		355		1 -		1	360	
129	ctc	ato	ttt	gca	caa	gac	ctc	atc	ctq	qac	agg	aat	gaq	qqa	qac	tgc	1338
						Asp											
131					365					370				•	375		
133	gt.g	gaa	ggc	atg	acq	gag	atc	ttc	gac	atg	ctg	ctg	gee	act	gct	tcc	1386
				_	_	Ğlu											
135			-	380					385					390			

DATE: 12/06/2001 PATENT APPLICATION: US/09/893,666 TIME: 10:36:14

Input Set : A:\210217US0.ST25.txt

137	cgc	ttc	cgt	gtg	ctc	aaa	ctc	aaa	cct	gag	gaa	ttc	gtc	tgc	ctc	aaa	1434
	Arg	Phe		Val	Leu	Lys	Leu	Lys	Pro	Glu	Glu	Phe	Val	Cys	Leu	Lys	
139			395					400					405				
1.41	gct	att	att	tta	ctc	aac	tcc	ggt	gct	ttt	tct	ttc	tgc	acc	ggc	acc	1482
142	Ala	Ile	Ile	Leu	Leu	Asn	Ser	Gly	Ala	Phe	Ser	Phe	Cys	Thr	Gly	Thr	
143		410					415					420					
	atg																1530
146	Met	Glu	Pro	Leu	His	Asn	Ser	Ala	Ala	Val	Gln	Ser	Met	Leu	Asp	Thr	
147	425					430					435					440	
149	atic	aca	gac	gca	ctc	att	cat	tac	atc	agt	cag	tcg	ggt	tac	ttg	qcc	1578
	Ile																
151					445					450			-	•	455		
153	cag	gag	cag	gcg	aga	cgg	cag	gcc	cag	ctg	ctc	ctg	ctg	ctc	tcc	cac	1626
	Gln																
155				460					465					470			
157	atic	agg	cac	atg	agc	aac	aaa	ggc	atq	gag	cac	ctc	tac	agc	atq	aaq	1674
	Ile																
159		-	475				-	480					485			1 -	
161	tgc	aag	aac	aaa	qtc	cct	ctt	tat	qac	ctc	cta	cta	gag	at.o	ctc	gat	1722
162	Cys	Lys	Asn	Lys	Val	Pro	Leu	Tyr	Ásp	Leu	Leu	Leu	Ğlu	Met.	Leu	Asp	
163	_	490		-			495	4	•			500					
165	gcc	cac	cqc	ctq	cac	cac	ccc	atc	aga	qca	ccc	caq	t.cc	tta	ticc	caa	1770
166	Ála	His	Arq	Leu	His	His	Pro	Val	Ara	Ala	Pro	Gln	Ser	Leu	Ser	Gln	1,,0
	505		,			510			)		515	0111		200	DCI	520	
169	gt.c	gac	aga	gac	cct.		tcc	acc	agc	agc		aaa	aat	gga	atc		1818
170	Val	Asp	Ara	Asp	Pro	Pro	Ser	Thr	Ser	Ser	Glv	Glv	Glv	Glv	Tle	Δla	10.10
171		1	)	L	525				001	530	OI1	O L J	OLI	Ory	535	nia	
	CCC	aat	tct	ata		αca	tct	саа	aac		atc	gag	agt	cca		ana	1866
174	Pro	Glv	Ser	Tle	Ser	Ala	Ser	Ara	Glv	Ara	Tle	Glu	Ser	Dro	Ser	Δra	1000
175		1		540			001	*** 9	545	111.9	110	GIU	DCI	550	DCI	Arg	
177	ggc	CCC	t.t.t.	-	aca	agt	atc	ctt		tat	gga	aaa	tca		cct	da c	1914
178	Gly	Pro	Phe	Ala	Pro	Ser	Val	Len	Gln	Tvr	Glv	Glv	Ser	Δra	Pro	Δen	17.14
179	. 1		555			001	, 41	560	01	- 1 -	O + I	OLI	565	111 9	110	АЗР	
181	tgc	acc		acc	att	саа	gac		icaca	aca o	rticca	aaac		++++	ttat		1965
182	Cys	Thr	Pro	Ala	Len	Gln	Asn	cyur	, caco	iou g	,	agge				-	1702
183	2	570			20 a	0.2.11	575										
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191	aσca	actt	†† †	ttta	ctct	c ca	aaca	acat	ata	ataa	att	ortaa	aatt	.ga g	rtaca	atgag	2205
																aagtg	2265
195	cact	t.cct	.ct. t	agat	taaa	ia an	agea	ator	i dea	ittat	+++	atet	ctta	rta o	acco	gggat.	2325
197	gact	.aaga	at a	atora	gagt	ਤ 99 a at	tato	acat	, yca : +++	atat	aga	actt	acco	iac e	aata	caatt	2385
199	ttaa	aata	ag a	ttta	aaaa	с аа	аапа	aaca	ara	teaa	acc	taaa	adce	ot a	aace aace	cacge	2445
201	tata	gaaa	ac t	.aaat	aaat	t to	ttat	CCAC	. aga	tato	tet	adaa	agra	ct +	tatt	ctctg	2505
203	tacc	t.aca	ac t	catt	tact	c ta	aact	tact	ant	tatt	gaa	catt	uyya tata	ca c	y - L . + + - + +	ccgtg	2565
205	tttt	tota.	arc a	ctat	aget	t at	gaac	acta	. чес	aada	atr.	taat	act+	ta a	taas	cagat	2625
	ttac	ctta	ta t	tata	cato	it ca	acca	caaa	cat	actt	tto	atac	gett	.cy a .a+ +	ract	gactg	2625 2685
209	cade	ttaa	tt a	aat+	atet	a aa	ant+	catt	taa	atat	tara.	atta	tata	yu t saan	yaut atta	gacty aaaag	2745
- 0 2		Legu	cc a	uucl	y cc c	u ua	uyıl	cutt	. caa	ucyc	cya	a L LC	LyLy	aa d	alld	aaaay	2/40

DATE: 12/06/2001 PATENT APPLICATION: US/09/893,666 TIME: 10:36:14

Input Set : A:\210217US0.ST25.txt

Output Set: N:\CRF3\12062001\I893666.raw

211 gcaattcctg tttctattt 2764 214 < 210> SEO ID NO: 2 215 <211> LENGTH: 575 216 <212> TYPE: PRT 217 · 213 > ORGANISM: Oryzias latipes 219 - 400> SEQUENCE: 2 221 Met Tyr Pro Glu Glu Ser Arg Gly Ser Gly Gly Val Ala Ala Val Asp 222 1 10 225 Phe Leu Glu Gly Thr Tyr Asp Tyr Ala Ala Pro Asn Pro Ala Thr Thr 229 Pro Leu Tyr Ser Gln Ser Ser Thr Gly Tyr Tyr Ser Ala Pro Leu Glu 3.5 4.0 233 Thr Asn Gly Pro Pro Ser Glu Gly Ser Leu Gln Ser Leu Gly Ser Gly 55 237 Pro Thr Ser Pro Leu Val Phe Val Pro Ser Ser Pro Arg Leu Ser Pro 70 75 241 Phe Met His Pro Pro Ser His His Tyr Leu Glu Thr Thr Ser Thr Pro 85 90 245 Val Tyr Arg Ser Ser His Gln Gly Ala Ser Arg Glu Asp Gln Cys Gly 105 249 Ser Arg Glu Asp Thr Cys Ser Leu Gly Glu Leu Gly Ala Gly Ala Gly 115 120 253 Ala Gly Gly Phe Glu Met Ala Lys Asp Thr Arg Phe Cys Ala Val Cys 254 135 140 257 Ser Asp Tyr Ala Ser Gly Tyr His Tyr Gly Val Trp Ser Cys Glu Gly 150 155 261 Cys Lys Ala Phe Phe Lys Arg Ser Ile Gln Gly His Asn Asp Tyr Met 165 170 265 Gys  $\operatorname{Pro}$  Ala Thr Asn Gln Gys Thr Ile Asp  $\operatorname{Arg}$  Asn  $\operatorname{Arg}$   $\operatorname{Arg}$  Lys  $\operatorname{Ser}$ 185 190 269 Cys Gln Ala Cys Arg Leu Arg Lys Cys Tyr Glu Val Gly Met Met Lys 195 200 205 273 Gly Gly Val Arg Lys Asp Arg Ile Arg Ile Leu Arg Arg Asp Lys Arg 210 215 220 277 Arg Thr Gly Val Gly Asp Gly Asp Lys Val Val Lys Gly Gln Glu His 230 235 281 Lys Thr Val His Tyr Asp Gly Arg Lys Arg Ser Ser Thr Gly Gly Gly 245 250 285 Gly Gly Gly Gly Gly Arg Leu Ser Val Thr Ser Ile Pro Pro Glu 265 270 289 Gln Val Leu Leu Leu Gln Gly Ala Glu Pro Pro Ile Leu Cys Ser 275 280 293 Arg Gln Lys Leu Ser Arg Pro Tyr Thr Glu Val Thr Met Met Thr Leu 290 295 297 Leu Thr Ser Met Ala Asp Lys Glu Leu Val His Met Ile Ala Trp Ala 298 305 310 315 301 Lys Lys Leu Pro Gly Phe Leu Gln Leu Ser Leu His Asp Gln Val Leu 302 325 330 305 Leu Leu Glu Ser Ser Trp Leu Glu Val Leu Met Ile Gly Leu Ile Trp

PATENT APPLICATION: US/09/893,666

DATE: 12/06/2001 TIME: 10:36:14

Input Set : A:\210217US0.ST25.txt

306 340 345	350
309 Arg Ser Ile His Cys Pro Gly Lys Leu Ile Phe Ala Glr	Asp Leu Ile
310 355 360 365	
313 Leu Asp Arg Asn Glu Gly Asp Cys Val Glu Gly Met Thr	Glu Ile Phe
314 370 375 380	
317 Asp Met Leu Leu Ala Thr Ala Ser Arg Phe Arg Val Leu	ı Lys Leu Lys
318 385 390 395	400
321 Pro Glu Glu Phe Val Cys Leu Lys Ala Ile Ile Leu Leu	Asn Ser Gly
322 405 410	415
325 Ala Phe Ser Phe Cys Thr Gly Thr Met Glu Pro Leu His	Asn Ser Ala
326 420 425	430
329 Ala Val Gln Ser Met Leu Asp Thr Ile Thr Asp Ala Leu	lle His Tvr
330 435 440 445	
333 Ile Ser Gln Ser Gly Tyr Leu Ala Gln Glu Gln Ala Arg	Arg Gln Ala
334 450 455 460	,
337 Gln Leu Leu Leu Leu Ser His Ile Arg His Met Ser	Asn Lvs Glv
338 465 470 475	480
341 Met Glu His Leu Tyr Ser Met Lys Cys Lys Asn Lys Val	
342 485 490	495
345 Asp Leu Leu Glu Met Leu Asp Ala His Arg Leu His	
346 500 505	510
349 Arg Ala Pro Gln Ser Leu Ser Gln Val Asp Arg Asp Pro	
350 515 520 525	
353 Ser Ser Gly Gly Gly Gly Ile Ala Pro Gly Ser Ile Ser	
354 530 535 540	ma ber mig
357 Gly Arg Ile Glu Ser Pro Ser Arg Gly Pro Phe Ala Pro	Ser Val Leu
358 545 550 555	560
361 Gln Tyr Gly Gly Ser Arg Pro Asp Cys Thr Pro Ala Leu	
362 565 570	575
365 <210 > SEQ ID NO: 3	3,3
366 <211> LENGTH: 20	
367 <212> TYPE: DNA	
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371 -: 223> OTHER INFORMATION: SYNTHETIC DNA	
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374 toggtgacat gtaccotgaa	20
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386 ctgtqtgctc agtcttgaag	20
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390 <211> LENGTH: 20	
391 <212: TYPE: DNA	
392 <213> ORGANISM: ARTIFICIAL SEQUENCE	
2	

09/893,666 6

<210> 6
<211> 20
<212> DNA
<213> ARTIFICIAL DNA

See Jem 10 on Even Jummary Sheet
<400> 6
gtaggaggtc ataaagaggg

### VERIFICATION SUMMARY

PATENT APPLICATION: US/09/893,666

DATE: 12/06/2001 TIME: 10:36:15

Input Set : A:\210217US0.ST25.txt

Output Set: N:\CRF3\12062001\1893666.raw

L:404 M:220 C: Keyword misspelled or invalid format, <213> ORGANISM for SEQ ID#:6

L:406 M:258 W: Mandatory Feature missing, <220> FEATURE: L:406 M:258 W: Mandatory Feature missing, <223> OTHER INFORMATION: